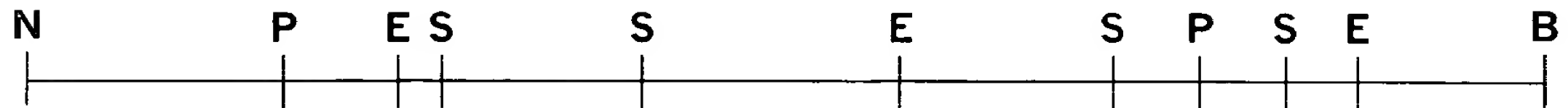


Figure 1: Restriction pattern of the HAL coding region cut with selected enzymes.

HAL



N - NdeI site introduced at the N-terminus

B - BamHI site introduced at the C-terminus

E - EagI

P - PstI

S - SphI

Figure 2: Experimentally derived peptide sequences of HAL

N-terminal

(M)ASAPQITLGLSGATAD

Internal

(M)ALADLDELLDEA

(M)GEPVEREVLRA

096374E-0410

Figure 3: SphI digestion pattern of HAL gene showing oligonucleotide and subclones.

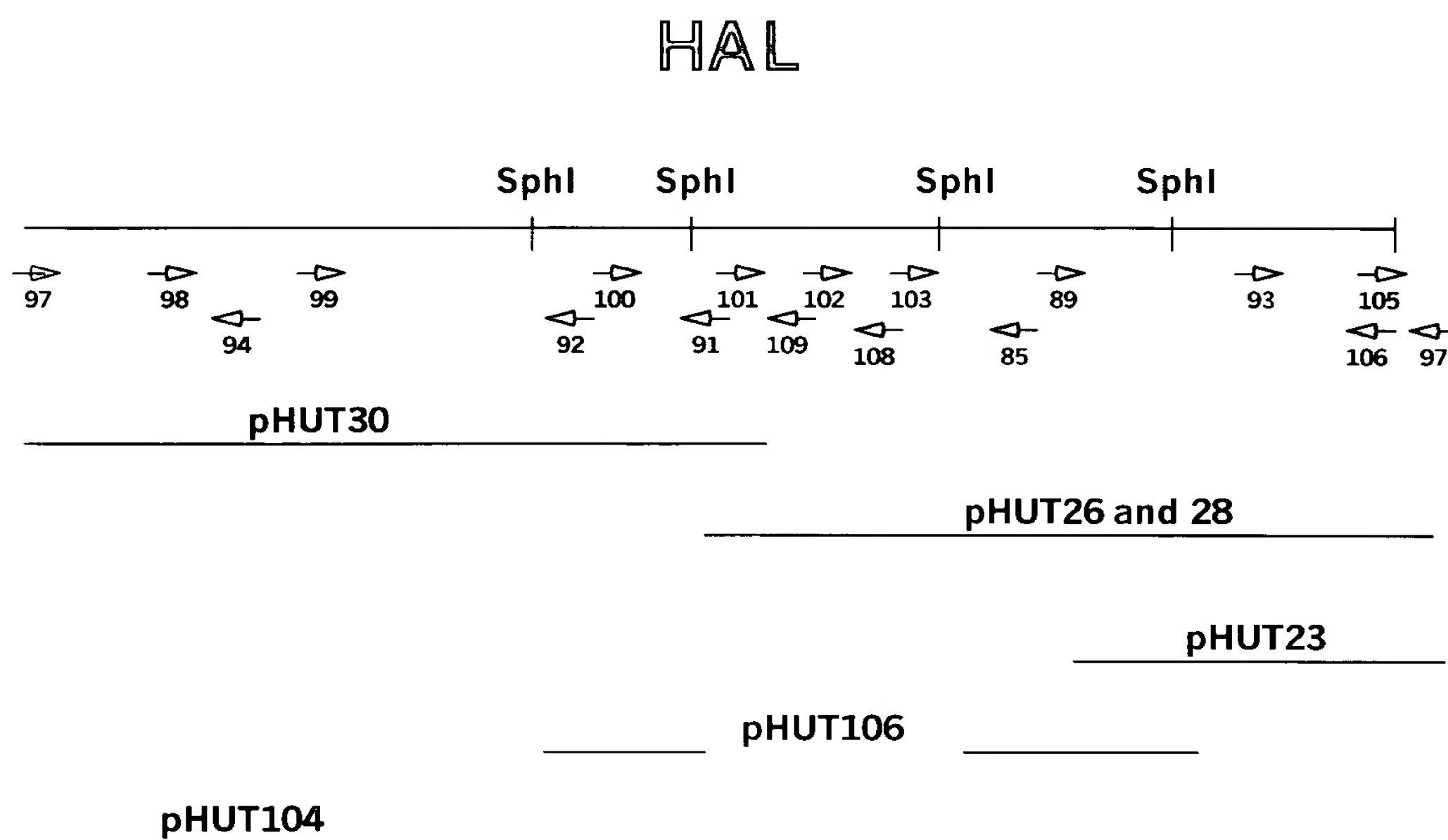
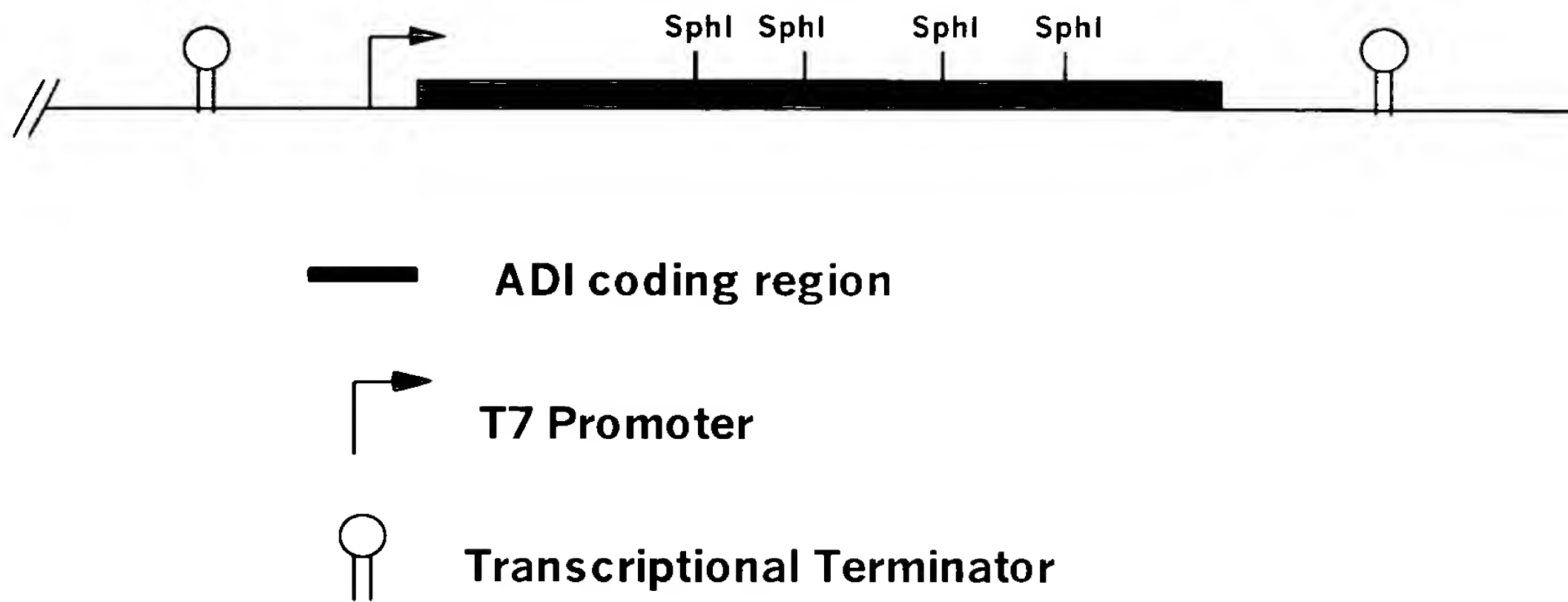


Figure 4: Histidine ammonia lyase overexpressing plasmid.

pHUT102



09833745-044304
10E740 5422260

Figure 5: SDS-PAGE showing expression of HAL in *E. coli*.

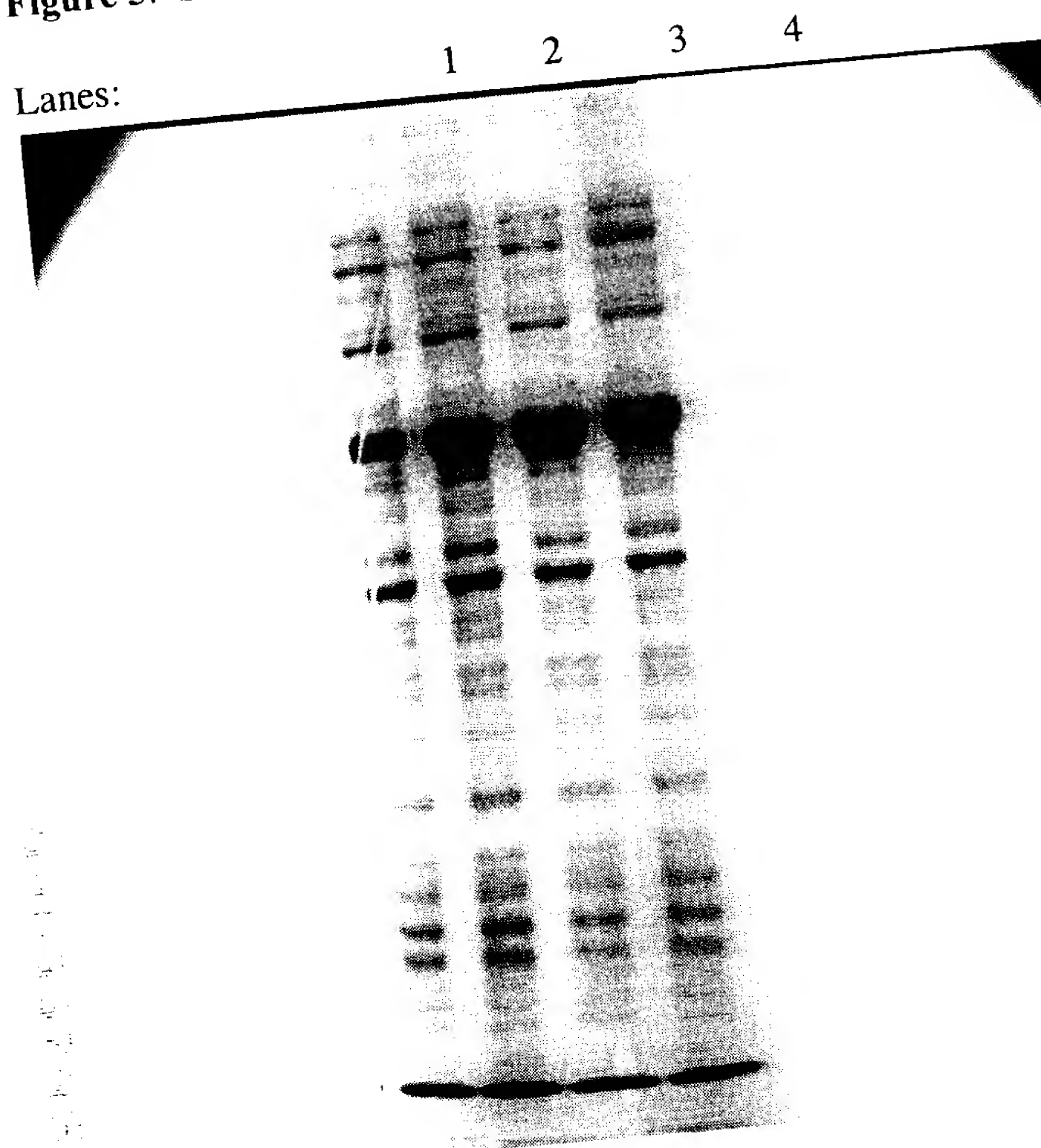
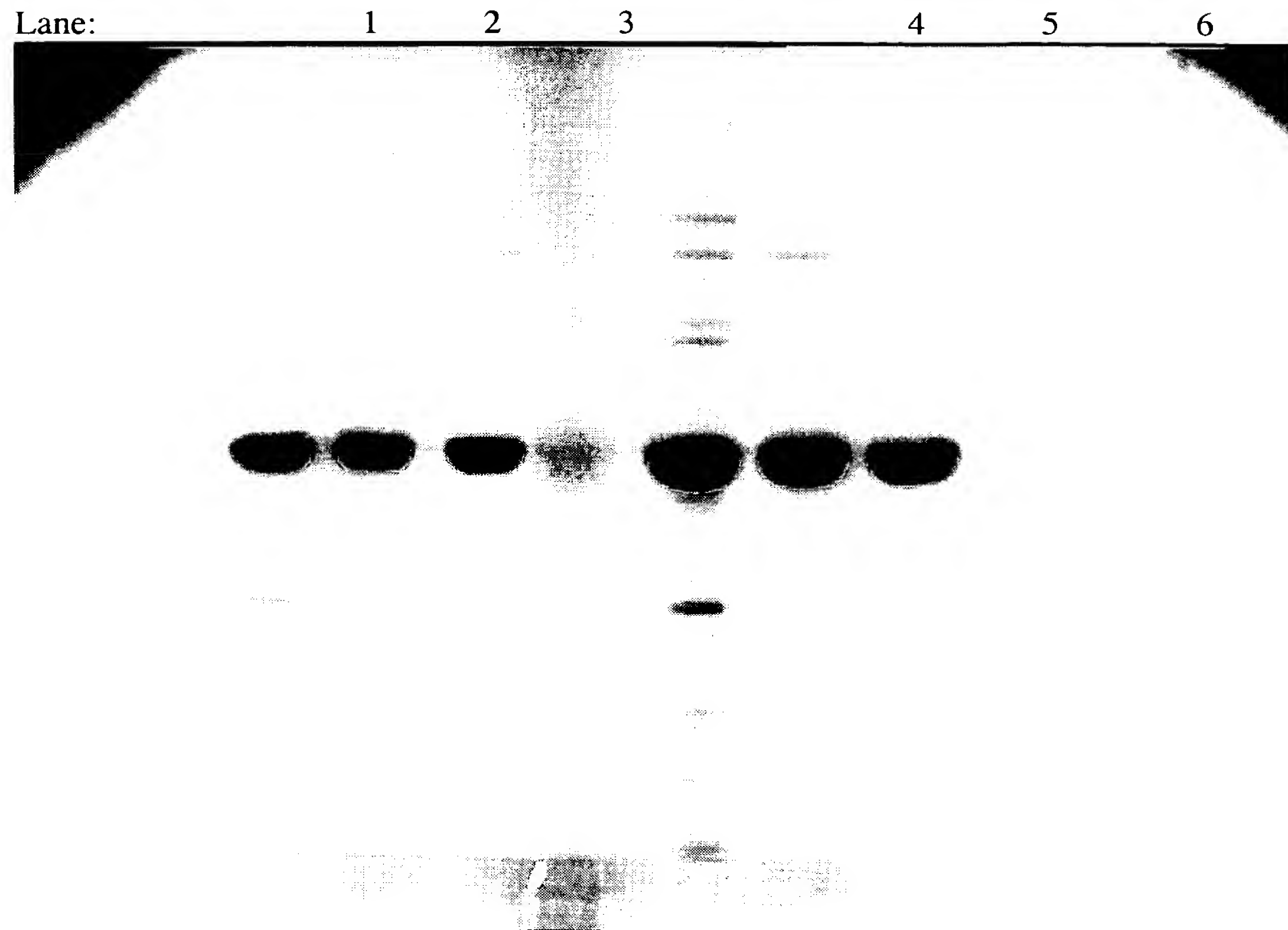


Figure 6: SDS-PAGE showing purification of HAL from *E. coli*



093745-044304

Figure 7: Effect of Temperature on HAL

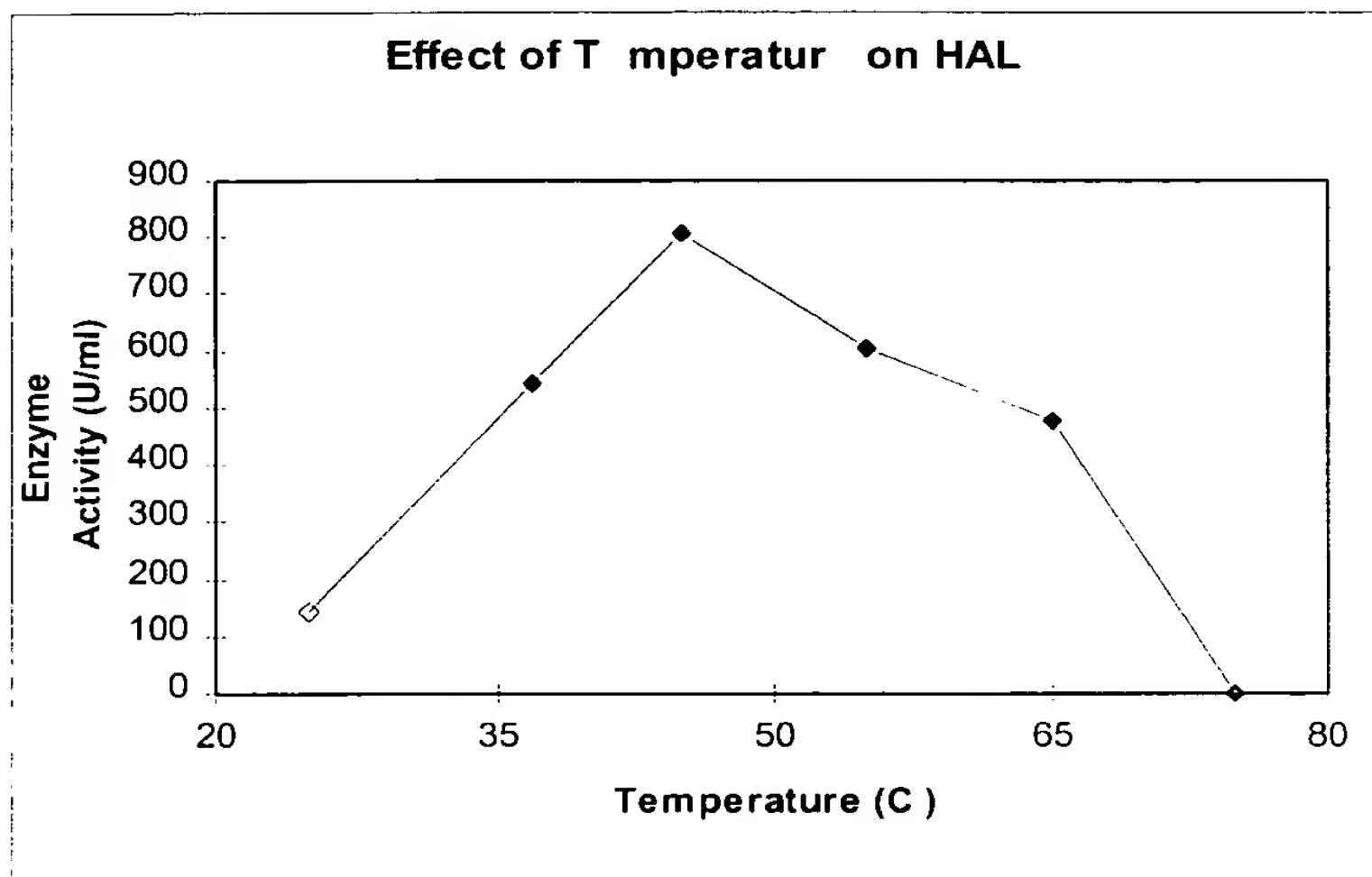
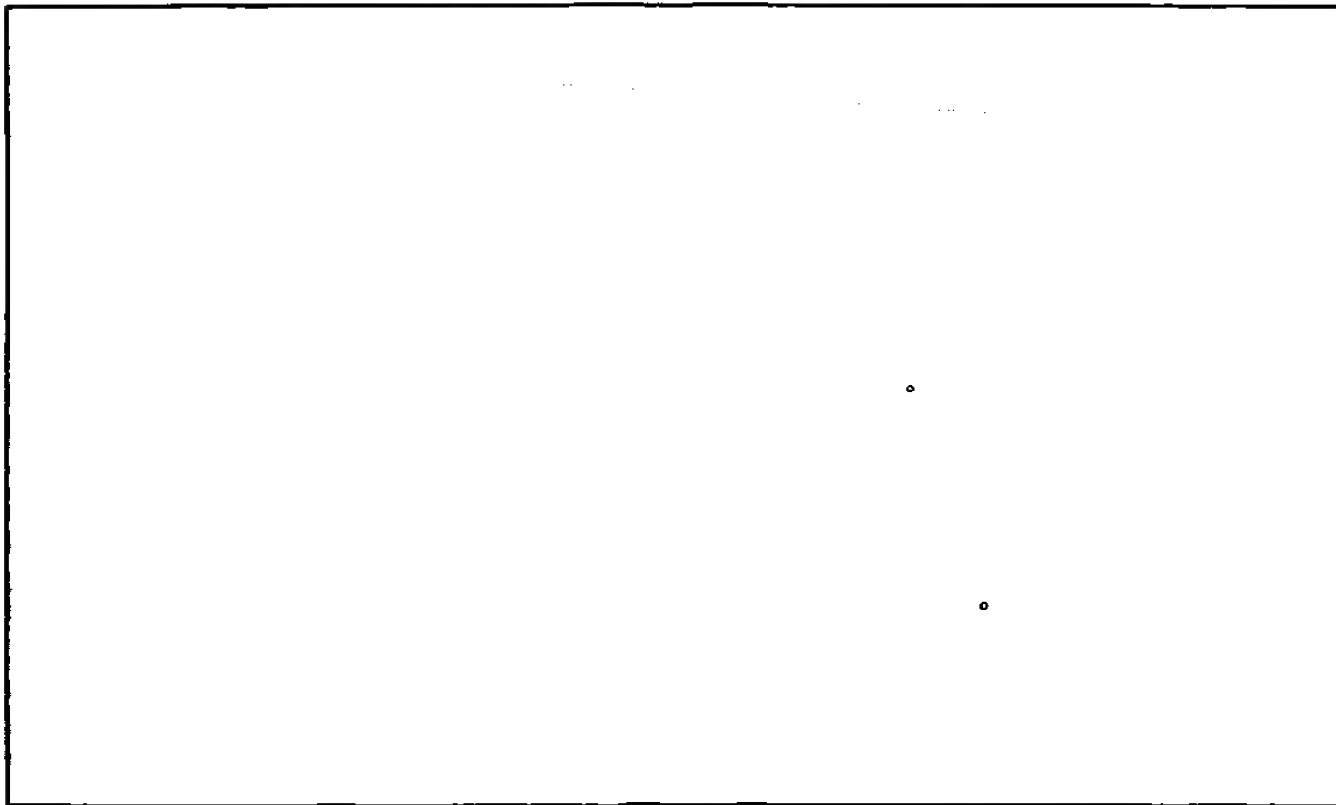


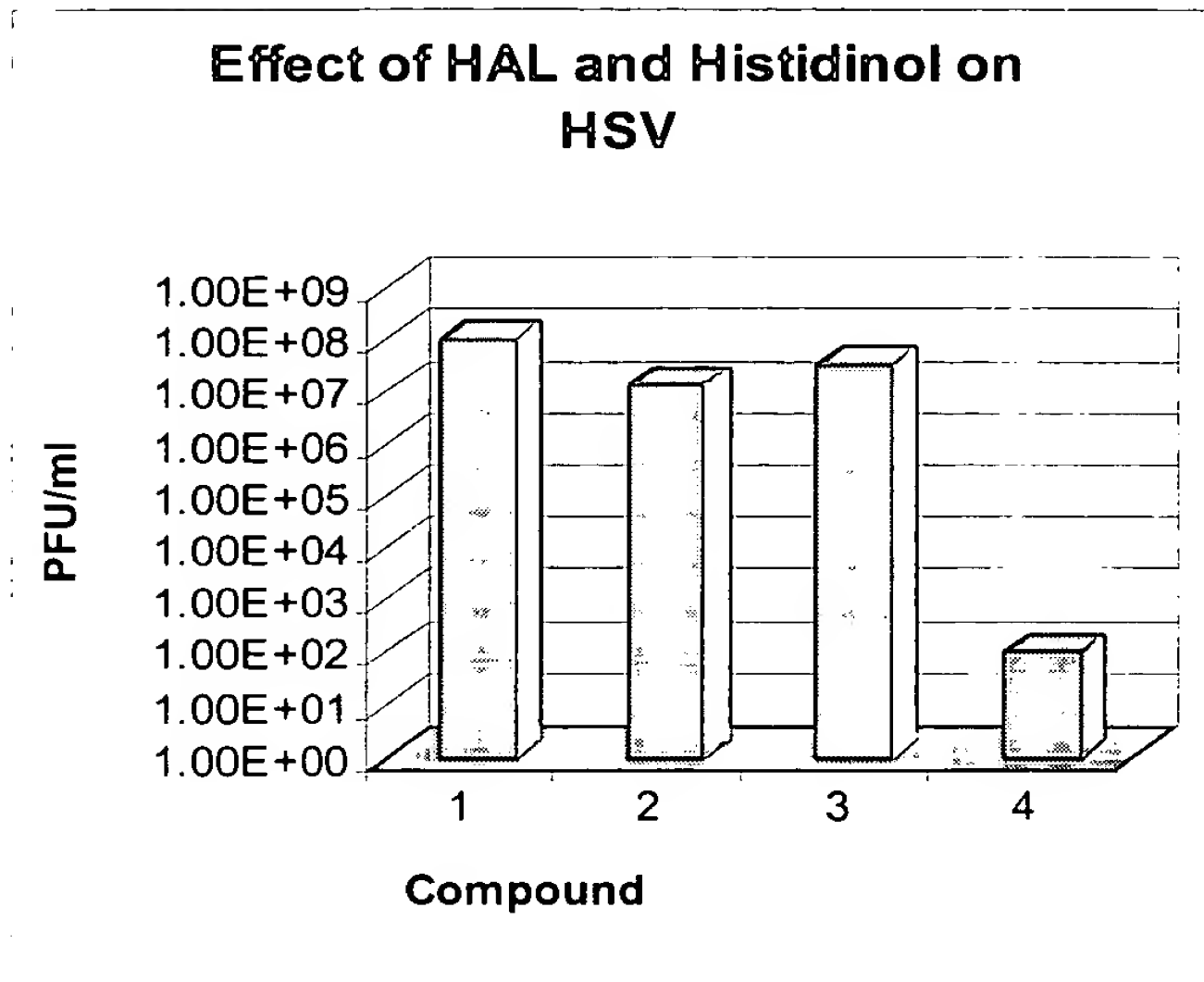
FIG. 7

Figure 8: Effect of pH on HAL.



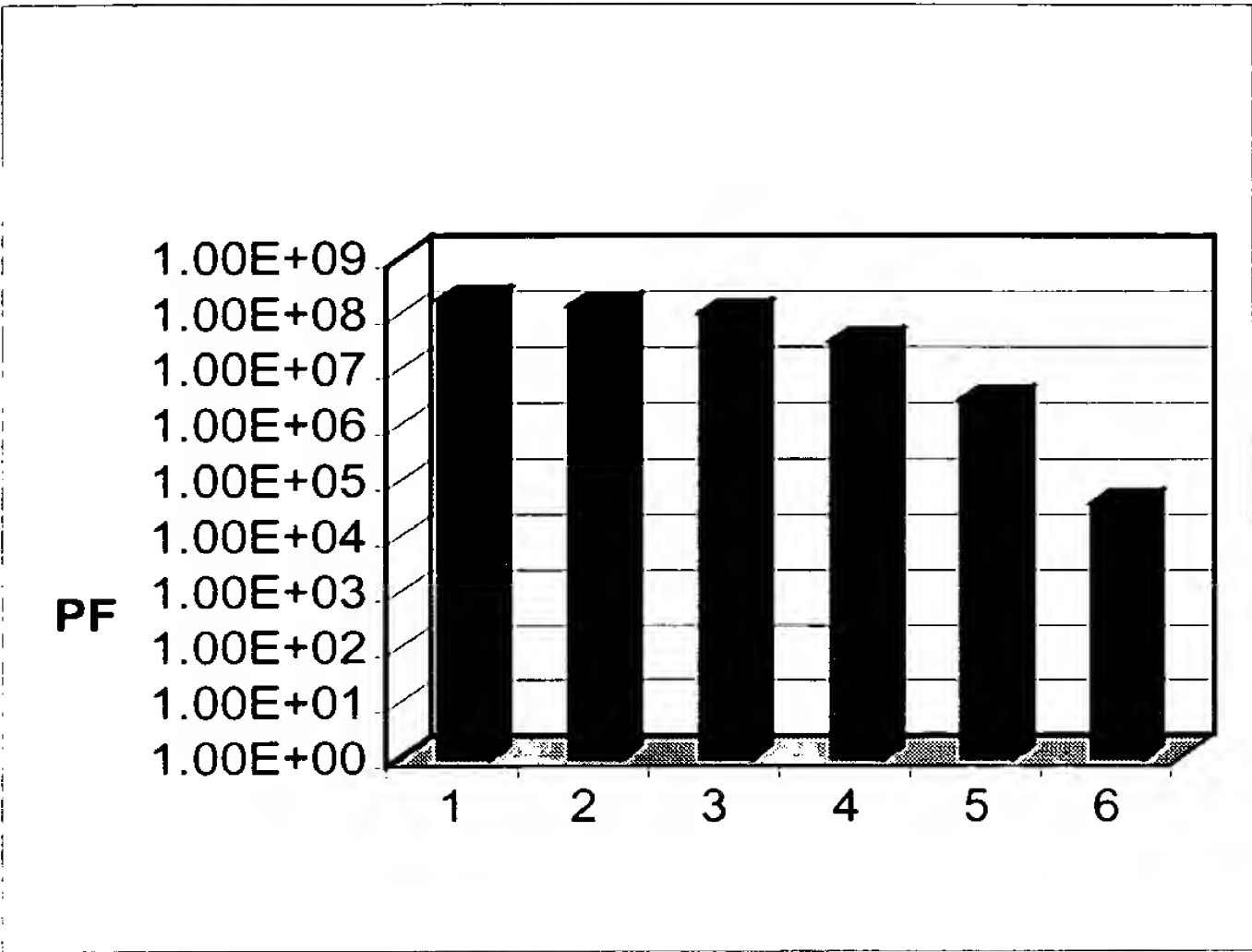
078728-0106

Figure 9: Effect of HAL and Histidinol on HSV.



078728-0106

Figure 10: Effectiveness of L-histidinol as a Single Agent



09032745-041304
00000000-00000000

Figure 11: Effect of HAL and Histidinol on RSV.

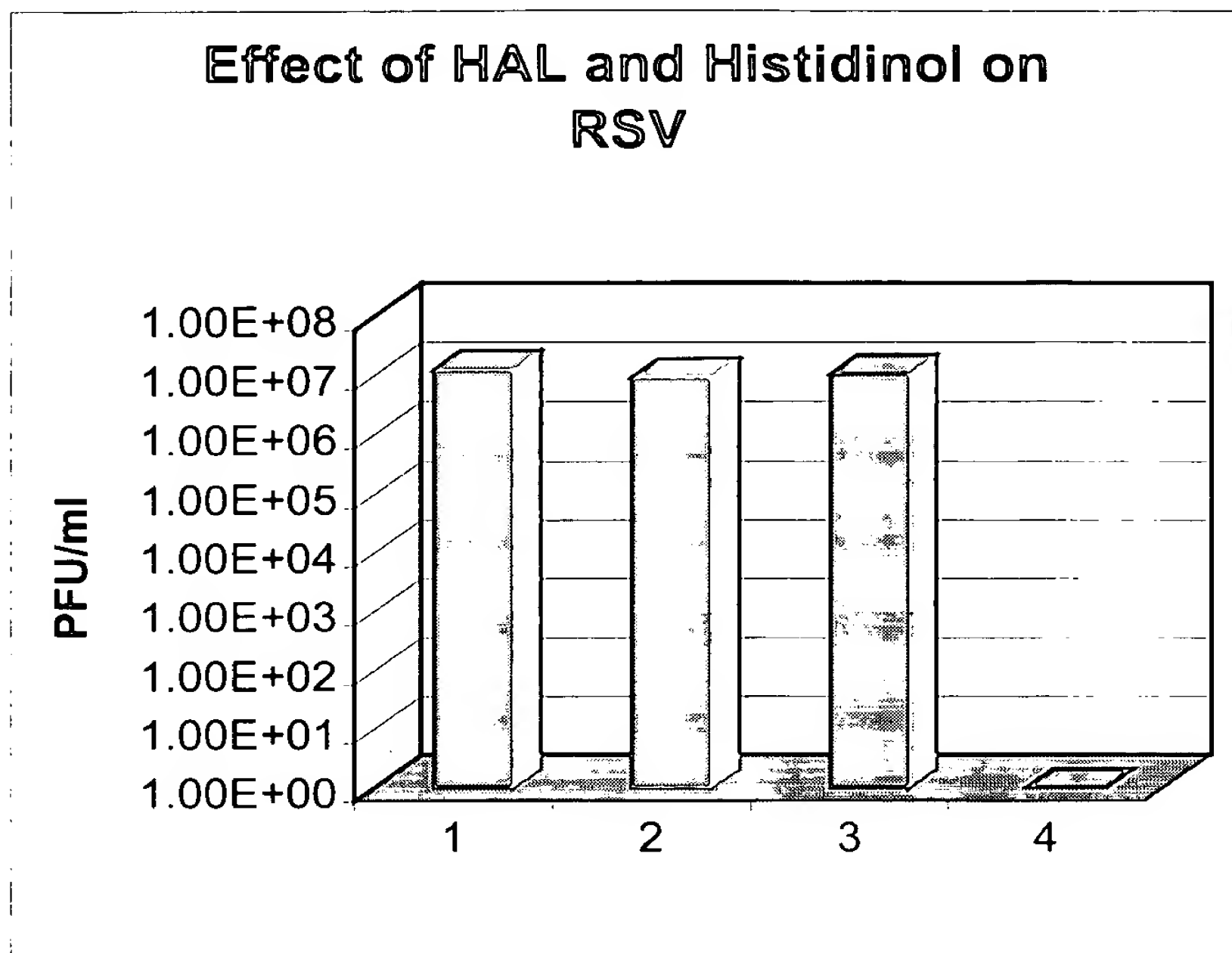
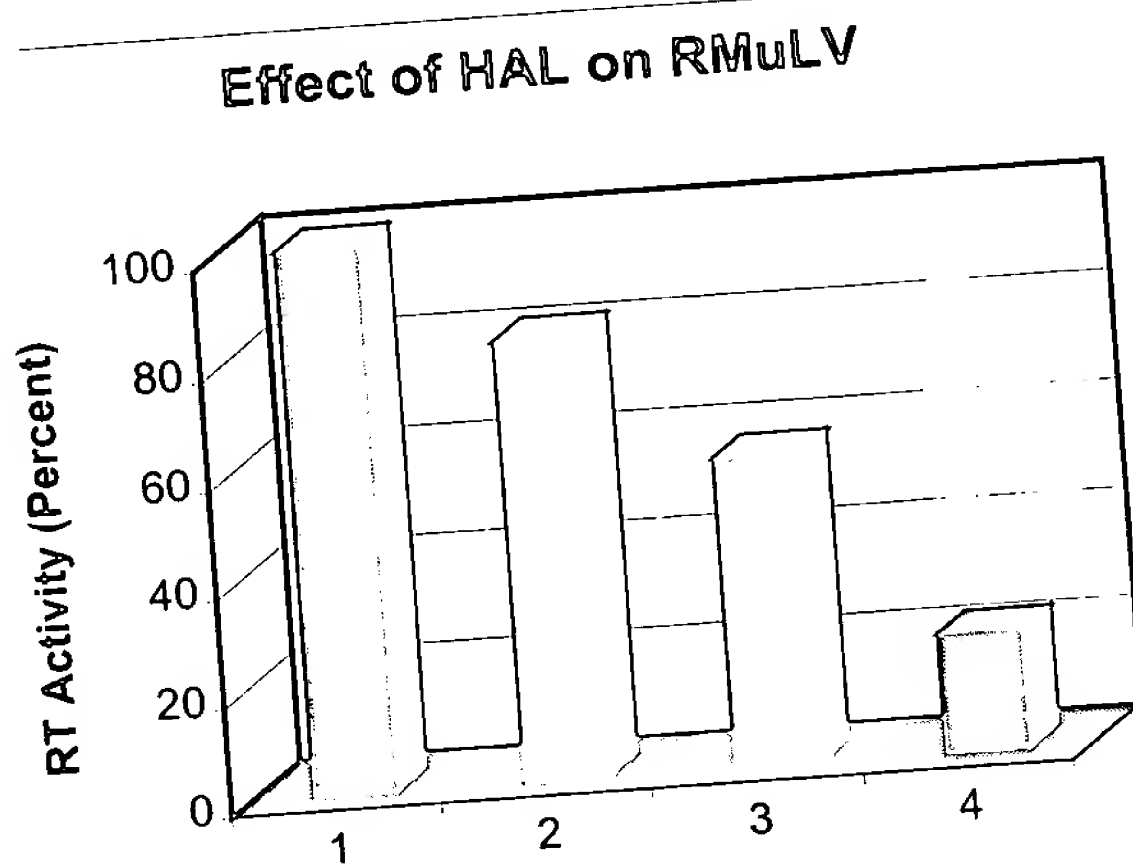


FIG. 11

Figure 12: Effect of HAL on RMuLV.



078728/0106

Figure 13: Histidine ammonia lyase peptide sequence pileup

```
HUTH_PSEPU -----
-
HUTH_RHIME -----
-
HUTH_MOUSE
MPRYTVHVRGEWLAVPCQDGKLTVGWLGREAVRRYMKNKPDNGGFTSVDEVQFLVHRCKG
HUTH_RAT
MPRYTVHVRGEWLAVPCQDGKLSVGWLGREAVRRYMKNKPDNGGFTSVDEVRFVRRCKG
HUTH_HUMAN
MPRYTVHVRGEWLAVPCQDAQLTVGWLGREAVRRYIKNKPDNGGFTSVDDAHFLVRRCKG
HUTH_CAEEL -MRLQVQIGTECVVVPCKP-DDTIHAVAKKSVEKLRLRLRPK----
LPLADDYFEVRRTVG
HUTH_BACS -----
-
HUTH_STRGR -----
-
HUTH_CORY -----
-

HUTH_PSEPU -----
-
HUTH_RHIME -----
-
HUTH_MOUSE LGLLDNEDELEVALEDNEFVEVVIEGDVMS-----PDFIPSQPEGVFLYSKYR---
-
HUTH_RAT LGLLDNEDLLEVALEDNEFVEVVIEGDVMS-----PDFIPSQPEGVFLYSKYR---
-
HUTH_HUMAN LGLLDNEDRLEVALENNEFVEVVIEGDAMS-----PDFIPSQPEGVYLYSKYR---
-
HUTH_CAEEL
NSLLDPEDLVSDVLKDSDFIIVAASVEETEDAKEAKKQEEIDNARAEIEKIDNRRRKVSF
HUTH_BACS -----
-
HUTH_STRGR -----
-
HUTH_CORY -----
-

HUTH_PSEPU -----
TELT LKPGT LTLAQLRAIHAAPVRLQLDASAAPAI DASVACVEQIIA
HUTH_RHIME -----
MTVI LRPGSVPLSDLETIYWTGAPARLDAAFDAGIAKAAARIAEIVA
HUTH_MOUSE -----
EPEKYIALDGD SLSTEDLVNLGKGGRYKIKLTSIAEKKVQQSREVIDSIK
HUTH_RAT -----
EPEKYIALDGD SLSTEDLVNLGKGGRYKIKLTSIAEKKVQQSREVIDSIK
HUTH_HUMAN -----
EPEKYI EL DGDRLTTEDLVNLGKGGRYKIKLTPTAEKRVQKSREVIDSIK
HUTH_CAEEL
ADSLAPMVLAPPTKLLILDGNSLLPEDLVRCEKGECAIQLSMESEDRIRKARTFLEKIAS
HUTH_BACS -----
MVTLDGSSLTTADVARVLFDFEEAAASEESMERVKKSRAAVERIVR
HUTH_STRGR -----
MDMHTVVVGTS GTTAEDVVAVARHGARVELSAAAVEALAAARLIVDALAA
HUTH_CORY -----
MASAPQITLGLSGATADDVIAVARHEARISISPQVLEELASVRAHIDALAS
```

Title: CLONING, OVEREXPRESSION AND
THERAPEUTIC USE OF BIOACTIVE
HISTIDINE AMMONIA LYASE
Inventor(s): Joseph ROBERTS et al.
DOCKET NO.: 078728/0106

Figure 13 cont'd.

Figure 13 cont'd.

HUTH_PSEPU
EDRTAYGINTGFGLLASTRIASHDLENLQORSLVLSHAAGIGAPLDDDLVRLIMVLKINSL
HUTH_RHIME
GNAPVYGINTGFGKGLASIKIDSSDVATLQORNILSHCCGVGQPLTEDIVRLIMALKLISL
HUTH_MOUSE
ERTVVYGITTGFGKFARTVIPANKLQELQVNLVRSHSSGVGKPLSPERCRMLLALRINVL
HUTH_RAT
ERTVVYGITTGFGKFARTVIPANKLQELQVNLVRSHSSGVGKPLSPERCRMLLALRINVL
HUTH_HUMAN
EKTVVYGITTGFGKFARTVIPINKLQELQVNLVRSHSSGVGKPLSPERCRMLLALRINVL
HUTH_CAEEL
EHRAVYGVTGFGTFSNVTIPPEKLKKLQNLIRSHATGYGEPLAPNRARMMLLALRINIL
HUTH_BACS
DEKTIYGINTGFGKFSDVLIQKEDSAALQNLILSHACGVGDPFPECVSRAMLLLRANAL
HUTH_STRGR
KPEPVYGVTGFGALASRHIGTELRAQLQORNIVRSHAAGMGPRVEREVVRALMFLRLKTV
HUTH_CORY
ADTPVYGISTGFGALATRHIAPEDRAKLQORSLIRSHAAGMGEPVEREVVRALMFLRAKTL

FOE T40 " 344E960

HUTH_PSEPU
SRGFSGIRRKVIDALIALVNAEVYPHIPLKGSVGASGDLAPLATMSLVLLGEGKARYKGO
HUTH_RHIME
GRGASGVRLELVRLIEAMLDKGVLIPIPEKGSVGASGDLAPLAHMAAVMMGHGEAFFAGE
HUTH_MOUSE
AKGYSGISLETLKQVIEAFNASCLSYVPEKGTVGASGDLAPLSHLALGLIGEGKMWSPKS
HUTH_RAT
AKGYSGISLETLKQVIEVFNASCLSYVPEKGTVGASGDLAPLSHLALGLIGEGKMWSPKS
HUTH_HUMAN
AKGYSGISLETLKQVIEMFNASCLPYVPEKGTVGASGDLAPLSHLALGLVGEKGMWSPKS
HUTH_CAEL
AKGHSGISVENIKMIAAFNAFCVSYVPQOGTVGCSGDLCPAHHLALGLLGEKGMWSPTT
HUTH_BACS
LKGFSGVRAELIEQLLAFLNKRVPVPIPOQGS LGASGDLAPLSHLALALIGQGEVFFEGE
HUTH_STRGR
ASGHTGVRPEVAQTMAVDNLNAGITPVVHEYGSLGCSGDLAPLSHCALTLMGEGEAEAGPDG
HUTH_CORY ASGRS-
VRPVVLETMVGMLNAGITPVVREYGSLGCSGDLAPLSHCALVLMGEGEATDAHG

REFVEEET
HUTH_PSEPU -
WLSATEALAVAGLEPLTLAAKEGLALLNGTQASTAYALRGLFYAEDLYAAAIACGGLSV
HUTH_RHIME -
RMKGDAALKAAGLSPVTLAAKEGLALINGTQVSTALALAGLFRAHRAGQAALITGALST
HUTH_MOUSE
GWADAKYVLEAHGLKPIVLKPKEGLALINGTQMITSLGCEALERASAIARQADIVAALT
HUTH_RAT
GWADAKYVLEAHGLKPIVLKPKEGLALINGTQMITSLGCEAVERASAIARQADIVAALT
HUTH_HUMAN
GWADAKYVLEAHGLKPVILKPKEGLALINGTQMITSLGCEAVERASAIARQADIVAALT
HUTH_CAEEL
GWQPADVVLLKKNLEPLELGPKEGLALINGTQMVTALGAYTLERAHNIARQADVIAALS
HUTH_BACS -
RMPAMTGLKKAGIQPVTLSKEGLALINGTQAMTAMGVVAYIEAEKLAYQTERIASLTI
HUTH_STRGR
TVRPAGELLAAHGIAPVELREKEGLALLNGTDGMLGMLVMALADLRNLYTSADITAALS
HUTH_CORY
DIRPVPELFAEAGLTPVELAEKEGLALVNGTDGMLGQLIMALADLDELDDIADATAAMSV

Figure 13 cont'd.

HUTH_PSEPU EAVLGSRSPFDARIHE-ARGQRGQIDTAACFRDLLGDSSEVSLSHKNCD----
KVQDPYS
HUTH_RHIME DAAMGSSAPFHPDIQH-CAAIRARSTRAAALRQLLTG-SPIRQSHIEGDE---
RVQDPYC
HUTH_MOUSE EVLKGTTKAFDTDIHA-VRPHRGQIEVAFRFRSL LDS-
DHHPSEIAESHRFCDRVQDAYT
HUTH_RAT EVLKGTTKAFDTDIHA-VRPHRGQIEVAFRFRSL LDS-
DHHPSEIAESHRFCDRVQDAYT
HUTH_HUMAN EVLKGTTKAFDTDIHA-LRPHRGQIEVAFRFRSL LDS-
DHHPSEIAESHRFCDRVQDAYT
HUTH_CAEEL DVLKGTTRAYDPDIHR-IRPHRGQNLSALRLRALLHS-
EANPSQIAESHRNCTKVQDAYT
HUTH_BACS EGLQGIIDAFDEDIHL-ARGYQEQIDVAERIRFYLS D-SGLTTSQGE-----
LRVQDAYS
HUTH_STRGR EALLGTDKVLAPELHA-IRPHPGQGV SADNMSRVLAG-SGLTGHHQDDAP---
RVQDAYS
HUTH_CORY EAQLGTDQVFRAELHEPLRPHPGQGRSAQNMFAFLAD-SPIVASHREGDG---
RVQDAYS

HUTH_PSEPU
LRCQPQVMGACLTQLRQAAEVLGIEANAVSDNPLVFAAEGDVISGGNFHAEPVAMAADNL
HUTH_RHIME IRCQPQVDGACLDLLRSVAATLTIEANAVTDNPLVLSDN-
SVVSGGNFHAEPVAFAADQI
HUTH_MOUSE
LRCCPQVHG VVNDTIAFVKDIITTELNSATDNPMVFASRGETISGGNFHGEYPAKALDYL
HUTH_RAT
LRCCPQVHG VVNDTIAFVKDIITTELNSATDNPMVFASRGETISGGNFHGEYPAKALDYL
HUTH_HUMAN
LRCCPQVHG VVNDTIAFVKNIITTELNSATDNPMVFANRGETVSGGNFHGEYPAKALDYL
HUTH_CAEEL
LRCVPQVHG VVHDTIEFVREIITTEMNSATDNPLVFADREEIISGGNFHGEYPAKALDFL
HUTH_BACS
LRCIPQVHGATWQTLGYVKEKLEIEMNAATDNPLIFNDGDKVISGGNFHGQPIAFAMDFL
HUTH_STRGR VRCAPQVNGAGRDTLDHAALVAGRELASSVDNPVVL PDG-
RVESNGNFHGAPVAYVLDFL
HUTH_CORY LRCSPQVTGAARDTIAHARLVATRELAAAIDNPVVLPSG-
EVTSNGNFHGAPVAYVLDFL

HUTH_PSEPU ALAIAEIGSLSERRISLMMDKHMS-
QLPPFLVENGGVNSGFMIAQVTAAALASENKALSH
HUTH_RHIME
ALAVCEIGAISQRRIALLVDPALSLRLPAFLAKKPGLNSGLMIAEVTSAALMSENKQLSH
HUTH_MOUSE AIGVHELAAISERRIERLCNPSLS-
ELPAFLVAEGGLNSGFMIAHCTAAALVSESKALCH
HUTH_RAT AIGVHELAAISERRIERLCNPSLS-
ELPAFLVAEGGLNSGFMIAHCTAAALVSESKALCH
HUTH_HUMAN AIGIHELAAISERRIERLCNPSLS-
ELPAFLVAEGGLNSGFMIAHCTAAALVSENKALCH
HUTH_CAEEL AIAVAELAQMSERRLERLVNKELS-
GLPTFLT PDGGLNSGFMTVQLCAASLVSENKVLCH
HUTH_BACS KIAISELANIAERRIERLVNPQLN-
DLPPFLSPHPGLQSGAMIMQYAAASLVSENKTLAH
HUTH_STRGR
AIVAADLGSICERRTDRLLDKNRSHGLPPFLADDAGVDSGLMIAQYTQAALVSEMKRLAV
HUTH_CORY
AIAVADLGSIAERRTDRMLDPARSRDLPAFLADDPGVDSGMMIAQYTQAGLVAENKRLAV

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T0040-547660

Figure 13 cont'd.

```

HUTH_PSEPU      AKLEKARQALRSEVA-HYDRDRFFAPDIEKAVELLAKG---S-LTGLLPAGVLPSSL---
-
HUTH_RHIME      PELQKAAA VRGVSS-SIEEDRYMADDLKAAGDLVASG---R-LAAAVSAGILPKLEN-
HUTH_MOUSE      TPLEKVYDLVRSVVR-
PWIKDRFMAPDIEAAHRLLLDQKVWEVAAPYIEKYRMEHIPESR
HUTH_RAT        TPLEKVYDLVRSVVR-
PWIKDRFMAPDIEAAHRLLLDQKVWEVAAPYIEKYRMEHIPESR
HUTH_HUMAN      TPLEKVYDLVRSVVR-
PWIKDRFMAPDIEAAHRLLEQKVWEVAAPYIEKYRMEHIPESR
HUTH_CAEEL      APLHKIYQLVRSVAP-
PLNEDRYMKPEIDAVLEMIRENRIWEAVLPHLETLEAMEELDPD
HUTH_BACS       SYTKQLFQEMRKVVP-SIQQDRVFSYDIERLTDWLKK----ESLIPDHQNKELRGMNI-
HUTH_STRGR      PASEAVVAALRAAGAEGPGPDRFLAPDLAAADTFVREG---R-LVAAVEPVTGPLA---
-
HUTH_CORY       PATGAVLEVLRSKVA-GPGQDRFLSAELEAAYDLLANG---S-VHKALEAHLPE-----

```

```

HUTH_PSEPU      -----
HUTH_RHIME      -----
HUTH_MOUSE      PLSPTAFSLESRLRKN SATIPESDDL----
HUTH_RAT        PLSPTAFSLESRLRKN SATIPESDDL----
HUTH_HUMAN      PLSPTAFSLQFLHKKSTKIPES EDL----
HUTH_CAEL       ALRQFTKTPTGIVQDRSMIPISDDEESIE
HUTH_BACS       -----
HUTH_STRGR      -----
HUTH_CORY       -----

```


Figure 14

	983831	1	80
1	SWALL: CAC21618	100.0%	MASAPQITLGLSGATADDVIAVARHEARISISISQPVLLEELASVRAHIDALASADTPVYGISTGFGALATRHIAPEDRAKLQ
2	SWALL: HUTH_STRGR	66.1%	---MHTVVVGTSGVTASDVLAVARAGARIELSEEAVALAAARSVVDALAAKPDVPYGVSTGFGALATRHIISPELRGLQ
3	SWALL: HUTH_DEIRA	65.4%	-MDMHTVVVGTSGTTAEDVVAVARHGARVELSAAAVEALAAARLIVDALAAKPEPVYGVSTGFGALASRHIGTELRAQLQ
4	SWALL: BAB16159	46.8%	-----MILDRDLNLEQFISVVRHGEQVELSAAARERIARARTVIEQIVEGDTPIYGVNTGFGKFENVQIDRSQLAQLQ
5	SWALL: Q9KWE4	42.0%	-----VPLHHLADIYWNNGSAKLDPSPFDAAVLKGAAARIAEIAAGNAPVYGINTEGFGKLASIKIDAADLATLQ
6	SWALL: HUTH_BACSU	42.0%	-----VPLHHLADIYWNNGSAKLDPSPFDAAVLKGAAARIAEIAAGNAPVYGINTEGFGKLASIKIDAADLATLQ
7	SWALL: Q9KSQ4	40.4%	-----MVTLDGSSLTADVARVLFDFEEAAAASEESMERVKKSRAAVERIVRDEKTIYGINTEGFGKFSVDVLIQKEDSAALQ
8	SWALL: Q9HU85	42.2%	---MLHLMIKPGQLSLKQLRQVSRSPVLSLDPEAIPAIAESAQVVEQVISEGRTVYGINTEGFGLLANTKIAPODLETQLQ
9	SWALL: Q9KBE6	41.7%	---MSLHLKPGQLTLADLRQAYLAPVRLSLDPSADAPIAASVACVENIIAEGRTAYGINTEGFGLLASTRISPADLEKLQ
10	SWALL: HUTH_PSEPU	39.3%	--MTNLKLLDGRSLSLHDLHRIIYEGETVGASDESMKVKQSRKAVEQIVADEKIIYGITTTGFGKFSDFIDPDDVENLQ
11	SWALL: HUTH_RHIME	41.7%	---TELTLKPGTLTLAQLRAIHAAPVRLQLDASAAPAIDASVACVEQIIAEDRTAYGINTEGFGLLASTRIASHDLENLQ
12	SWALL: Q9HU90	40.6%	-----LRPGSVPLSDLETIYWTGAPARLDAAFDAGIAKAAARIAEIVAGNAPVYGINTEGFGKLASIKIDSSDVATLQ
13	SWALL: HUTH_HUMAN	40.7%	MSDLPSVVFGDGPLRWQELVAVARHGARLELSAAAWARIDNARAIVCRIVANGERAYGISTGLGALCDVLLGEQLAELS
14	SWALL: HUTH_CAEEL	39.2%	KYREPEKYIELDGLTTEDLVNLGKGRYKIKLTPTAEKRVQKSREVIDSIIKEKTVVYGITTTGFGKFA-RTVIPINKLQLQ
15	SWALL: Q9HLI6	38.8%	VLAPPTKLLILDGNSPEDLVRCCKECAIQLSMESEDRIRKARTFLEKIASSEHRAVYGVTTGFGTFSNVTIPPPEKLKKLQ
16	SWALL: HUTH_MOUSE	41.0%	-----MIEIDGRSLRVEDVYAVAVEYDRVSIISDDTLKAVEEKHEAFLKLINSGKTVYGVNTGFGSLLNVHIERDQEIELQ
17	SWALL: BAB29407	38.6%	KYREPEKYIALDGDSTEDLVNLGKGRYKIKLTSIAEKKVQQSREVIDSIIKERTVVYGITTTGFGKFA-RTVIPANKLQLQ
18	SWALL: HUTH_RAT	38.6%	KYREPEKYIALDGDSTEDLVNLGKGRYKIKLTSIAEKKVQQSREVIDSIIKERTVVYGITTTGFGKFA-RTVIPANKLQLQ
19	SWALL: AAG53586	38.2%	KYREPEKYIALDGDSTEDLVNLGKGRYKIKLTSIAEKKVQQSREVIDSIIKERTVVYGITTTGFGKFA-RTVIPANKLQLQ
20	SWALL: Q9KKE0	39.8%	---MNALTLPGTTLAQLRQVWQQPLQLTDESAHEAINDSVACVEAIVAEGRRTAYGINTEGFGLLAQTRIAATHDLENLQ
21	SWALL: Q9HQD5	38.9%	-----MGEMISLDGPLTWREIASIAEGASLDLSGPARRIAQARRIVDALVERGIRGYGINTEGFGALCDVIIISRENQQALS
		42.2%	-----MSDTRIDAADREALQ

Figure 14, cont'd.

	81	1	160
983831	100.0%	[RSLIRSHAAAGMGEPPERVVRRALMFLRAKTLASGRTGVRPVVLETMVGMNLNAGITPVVREYGSGLGCSGDLAPLSHCALVL
1 SWALL: CAC21618	66.1%	RNIVRSHAAAGMGEPPERVVRRALMFLRLKTVCSGRTGVRPEVAQTMADVLNAGITPVVHEYGSGLGCSGDLAPLSHCALT	TL
2 SWALL: HUTH_STRGR	65.4%	RNIVRSHAAAGMGEPPERVVRRALMFLRLKTVASGHTGVRPEVAQTMADVLNAGITPVVHEYGSGLGCSGDLAPLSHCALT	TL
3 SWALL: HUTH_DEIRA	46.8%	HNLIIVSHAIGMGEPLPAEVRVGMILLRAQSLSLGSHGVRVEVELLALLNADALPVVPSQGSVGASGDLAPLAHLALGL	
4 SWALL: BAB16159	42.0%	RNLIILSHCCGVGAPLPENNVRLIMALKLISLGRGASGVRIELIRLIEGMLEKGVIPVPEKGSVGASGDLAPLAHMSATM	
5 SWALL: Q9KWE4	42.0%	RNLIILSHCCGVGAPLPENNVRLIMALKLISLGRGASGVRIELIRLIEGMLEKGVIPVPEKGSVGASGDLAPLAHMSATM	
6 SWALL: HUTH_BACSU	40.4%	LNLIILSHACGVGDPFPECVSRAMLLLRANALLKFGSGVRAELIEQLLAFLNKRVRHPVIPPQQGSGLGASGDLAPLSHLALAL	
7 SWALL: Q9KSQ4	42.2%	KSIVLSHAAGIGELMSDETVRLMMLLKINSLARGYSGIRLEVIQALIELVNNQIYPCVPPKGSVGASGDLAPLAHMSVTL	
8 SWALL: Q9HU85	41.7%	RSIVLSHAAGVGEALDDAMVRLVMLLKVNLSLARGFSGIRRKVIDALIALINAEVYPHIPPLKGSVGASGDLAPLAHMSLVL	
9 SWALL: Q9KBE6	39.3%	HNLIYSHACGVSPFPETVSRMTLVLRANALLKFGSGVRPLVIERLLALVNANIHPVIPPQQGSGLGASGDLAPLSHLALVL	
10 SWALL: HUTH_PSEPU	41.7%	RSIVLSHAAGIGAPLDDDLVRLIMVLKINSLSRGFGIRRKVIDALIALVNAEVYPHIPPLKGSVGASGDLAPLAHMSLVL	
11 SWALL: HUTH_RHIME	40.6%	RNLIILSHCCGVGQPLTEDIVRLIMALKLISLGRGASGVRLVRLIEAMLDKGVIPVPEKGSVGASGDLAPLAHMAAVM	
12 SWALL: Q9HU90	40.7%	RNTLLSHACGVGEPLRDEQTRAIICAAVANYSQKSGLDRLSVEGLLALLNHGITPQVPAQGSVGY--LTHMAHVGI	AL
13 SWALL: HUTH_HUMAN	39.2%	VNLVRSHSSGVGKPLSPERCRLMLALRINVLAKYSGISLETLKQVIEAFNASCPLPYVPEKGTVGASGDLAPLSHLALGL	
14 SWALL: HUTH_CAEEL	38.8%	LNLIIRSHATGYGEPLAPNRARMLLALRINILAKHSGISVENIKMIAAFNAFCVSVYPQQGTVGCSGDLCPPLAHLALGL	
15 SWALL: Q9HLI6	41.0%	KNLIIRSHSSGVGDYLENRYVRAIMAVRLNSLAAGYSAVSADLLNMMVEMLNRDVIPA	VPKYGSVGASGDLAPLAHIGLAM
16 SWALL: HUTH_MOUSE	38.6%	VNLVRSHSSGVGKPLSPERCRLMLALRINVLAKYSGISLETLKQVIEAFNASC	LSYVPEKGTVGASGDLAPLSHLALGL
17 SWALL: BAB29407	38.6%	VNLVRSHSSGVGKPLSPERCRLMLALRINVLAKYSGISLETLKQVIEAFNASC	LSYVPEKGTVGASGDLAPLSHLALGL
18 SWALL: HUTH_RAT	38.2%	VNLVRSHSSGVGKPLSPERCRLMLALRINVLAKYSGISLETLKQVIEAFNASC	LSYVPEKGTVGASGDLAPLSHLALGL
19 SWALL: AAG53586	39.8%	RSIVLSHAAGVGEPLDDDIVRLMMVLKINSLARGFSGIRLSVIQALIALVNAGVSVDP	AKGSVGASGDLAPLAHMSLTL
20 SWALL: Q9KKE0	38.9%	RNIIILSHACGVGDPLGRVEARAVMAAQIANLTHGYSGVRVETAEMLLALLNADI	IPLIPSRGSVGY-----LTHAALVL
21 SWALL: Q9HQD5	42.2%	ANLVRSHAAAGAGSELDTAAVRALLVTRLNALAKYSGIRERVLDVLVGLLNEG	HPVVPSPRGSGLGASGDLAPLAHMSRVL

Figure 14, cont'd.

983831	100.0%	161	[2	240
1 SWALL: CAC21618	66.1%		MGE	EATDAHGDIRVPPELFAEAGLTPVELAEKEGLALVNGTDGMLGQIMALADLDELDDIADATAAMSVEAQLGTDQV	
2 SWALL: HUTH_STRGR	65.4%		MGE	DAEGPDGTVRPAGELLAAHGIAPVELREKEGLALLNGTDGMLGMLVMALADLTLYKSADITAAALTMEALLGTD	DRV
3 SWALL: HUTH_DEIRA	46.8%		MGE	EAEGPDGTVRPAGELLAAHGIAPVELREKEGLALLNGTDGMLGMLVMALADLRNLYTSADITAAALSLEALLGTD	DKV
4 SWALL: BAB16159	42.0%		I	GLDI-EYQGVQVRPAADVLAELGLSPVQLQAKEGLALINGTQLMGSLALALHDAQVLLGTANLAAAMTVEARYG	SHRP
5 SWALL: Q9KWE4	42.0%		MGE	GEAF-YQGVQMPKDALAKAGLSPVVLAKEGLALINGTQTSTALALAGLFRHRAAQSAALVTGALSTDAAMG	SSAP
6 SWALL: HUTH_BACSU	40.4%		MGE	GEAF-YQGVQMPKDALAKAGLSPVVLAKEGLALINGTQTSTALALAGLFRHRAAQSAALVTGALSTDAAMG	SSAP
7 SWALL: Q9KSQ4	42.2%		I	QGEVF-FEGERMPAMTGLKKAGIQPVTLTSKEGLALINGTQAMTAMGVVAYIEAEKLAYQTERIASLTIEGLQ	GIIDA
8 SWALL: Q9HU85	41.7%		LGE	GQAR-YNGKIIISGLEAMKIAGLEPITLAPKEGLALLNGTQASTAFALEGLFVAEDLFASATVCGAMSVEAALG	SRRP
9 SWALL: Q9KBE6	39.3%		I	GESRARH-RGEWLPAAEALAVAGLEPLTLAAKEGLALINGTQVSTAYALRGLFEAEDLFAAATVCGGLSVEAM	LGRAP
10 SWALL: HUTH_PSEPU	41.7%		LGE	GEVF-YKGTKTASFALKEEIEPITLTAKKEGLALINGTQAMTAMGVVAYIEAEKLAQSEIIASLTMEGLR	GIIDA
11 SWALL: HUTH_RHIME	40.6%		LGE	KAR-YKGWLSATEALAVAGLEPLTLAAKEGLALINGTQASTAYALRGLFYAEDLYAAAIACGGLSVEAVL	GSRSP
12 SWALL: Q9HU90	40.7%		M	GEAFFAGERMKGDAALKA-AGLSPVTLAAKEGLALINGTQVSTALALAGLFRHRAAGQAALITGALSTDAAMG	SSAP
13 SWALL: HUTH_HUMAN	39.2%		L	GIEVS-YRGSVVPAAAAAAEGLATVRLGAKDGLCLVNGTPCMTGLACLALDDAQRQAQWADVIGAMSFEALR	GQLAA
14 SWALL: HUTH_CAEEL	38.8%		V	GKMWSPKSGWADAKYVLEAHGKPVILKPKKEGLALINGTQMITSLGCEAVERASAIARQADIVAALTLEVLK	GTTKA
15 SWALL: Q9HLI6	41.0%		L	GKMWSPTTGWQPADVVILKNNLEPLELGPKEGLALINGTQMVVTALGAYTLERAHNIAHQADVIAALS	LDV
16 SWALL: HUTH_MOUSE	38.6%		M	GKAF-FEGRLMDSARALEKAGLKPYPQFKEKEGVALINGTSFMSGILSIAVMDAHDILENAIRSALLSFEAL	GGT
17 SWALL: BAB29407	38.6%		I	GKMWSPKSGWADAKYVLEAHGKPIVLKPKKEGLALINGTQMITSLGCEALERASAIARQADIVAALTLEVLK	GTTKA
18 SWALL: HUTH_RAT	38.2%		I	GKMWSPKSGWADAKYVLEAHGKPIVLKPKKEGLALINGTQMITSLGCEALERASAIARQADIVAALTLEVLK	GTTKA
19 SWALL: AAG53586	39.8%		L	GKAR-YRGEWLPAAATALQKAGLAPVTLAAKEGLALINGTQASTAFALRGLFEAEDLFASAVVCGALTTEAVL	GSRRP
20 SWALL: Q9KKE0	38.9%		I	HGSAMQGTERLSGADAL-ARLGLAPLRLEAKEGLSLVNGTPCATGLAALALARTERLFAWADAAAAAMTYE-NL	GSQAN
21 SWALL: Q9HQD5	42.2%		I	GQA-DVAGERMPAAEALAAADLEPVTLQAKEGLALINGTQLTGTGVAALALVDAERVLSADTAGALTTEVTM	STTAS

Figure 14, cont'd.

Title: CLONING, OVEREXPRESSION AND
THERAPEUTIC USE OF BIOACTIVE
HISTIDINE AMMONIA LYASE
Inventor(s): Joseph ROBERTS et al.
DOCKET NO. : 078728/0106

Figure 14, cont'd.

	983831	321	400
1	SWALL: CAC21618	100.0%	VVLP
2	SWALL: HUTH_STRGR	66.1%	PDGR
3	SWALL: HUTH_DEIRA	65.4%	PDGR
4	SWALL: BAB16159	46.8%	LIFPT
5	SWALL: Q9KWE4	42.0%	LVLS
6	SWALL: HUTH_BACSU	42.0%	LVLS
7	SWALL: Q9KSQ4	40.4%	LIFND
8	SWALL: Q9HU85	42.2%	LVFAD
9	SWALL: Q9KBE6	41.7%	LVFAA
10	SWALL: HUTH_PSEPU	39.3%	LIFDNG
11	SWALL: HUTH_RHIME	41.7%	LVFAAG
12	SWALL: Q9HU90	40.6%	LIFDNG
13	SWALL: HUTH_HUMAN	40.7%	LVFAAG
14	SWALL: HUTH_CAEEL	41.7%	LVLS
15	SWALL: Q9HLI6	40.7%	LLGT
16	SWALL: HUTH_MOUSE	39.2%	MVFAN
17	SWALL: BAB29407	38.8%	LVFAD
18	SWALL: HUTH_RAT	41.0%	L-FNG
19	SWALL: AAG53586	38.6%	MVFAS
20	SWALL: Q9KKE0	38.6%	MVFAS
21	SWALL: Q9HQD5	38.2%	MVFAS
		39.8%	LVFAA
		38.9%	AVAGS
		42.2%	LVFP

[illegible]

480

Figure 14, cont'd.

983831		100.0%	401
1 SWALL: CAC21618		66.1%	
2 SWALL: HUTH_STRGR		65.4%	
3 SWALL: HUTH_DEIRA		46.8%	
4 SWALL: BAB16159		42.0%	
5 SWALL: Q9KWE4		42.0%	
6 SWALL: HUTH_BACSU		40.4%	
7 SWALL: Q9KSQ4		42.2%	
8 SWALL: Q9HU85		41.7%	
9 SWALL: Q9KBE6		39.3%	
10 SWALL: HUTH_PSEPU		41.7%	
11 SWALL: HUTH_RHIME		40.6%	
12 SWALL: Q9HU90		40.7%	
13 SWALL: HUTH_HUMAN		39.2%	
14 SWALL: HUTH_CAEEL		38.8%	
15 SWALL: Q9HLI6		41.0%	
16 SWALL: HUTH_MOUSE		38.6%	
17 SWALL: BAB29407		38.6%	
18 SWALL: HUTH_RAT		38.2%	
19 SWALL: AAG53586		39.8%	
20 SWALL: Q9KKE0		38.9%	
21 SWALL: Q9HQD5		42.2%	

[illegible]

Figure 14, cont'd.

	983831	481		[.	5	.]	513
		100.0%		PGQDRFLSAELEAAYDLLANGSVHKALEHLPA					
1	SWALL:CAC21618	66.1%		PGPDRHLAPDLAAADAFVRAGHLVAAAEVSTGP					
2	SWALL:HUTH_STRGR	65.4%		PGPDRFLAPDLAAADTFVREGRLVAAVEPVTGP					
3	SWALL:HUTH_DEIRA	46.8%		LTEDRYFRPDLLRLRGELVSGRVAQAADTQAPA					
4	SWALL:BAB16159	42.0%		LEDDRYMATDLKAAIEVVASGALVSAISSGLPV					
5	SWALL:Q9KWE4	42.0%		LEDDRYMATDLKAAIEVVASGALVSAISSGLPV					
6	SWALL:HUTH_BACSU	40.4%		IQQDRVFSYDIERLTDWLKKESLIPDHQNKELR					
7	SWALL:Q9KSQ4	42.2%		YDKDRYFAPDIEKANALL-QLAVHNRLMPDQLL					
8	SWALL:Q9HU85	41.7%		YQEDRFFAPDIEAASQLLASGCLNALLPARLLP					
9	SWALL:Q9KBE6	39.3%		IDQDRMFAKDIERAAKWLDGSDFTKMREKER					
10	SWALL:HUTH_PSEPU	41.7%		YDRDRFFAPDIEKAVELLAKGSLTGLLPAGLPS					
11	SWALL:HUTH_RHIME	40.6%		IEEDRYMADDLKAAGDLVASGRLAAAVSAGLPK					
12	SWALL:Q9HU90	40.7%		YDTRWLAPDIAASAAI LGERKSLARLAASIGD					
13	SWALL:HUTH_HUMAN	39.2%		WIKDRFMAPDIEAAHRLLLLEQKVWEVAAPYIEK					
14	SWALL:HUTH_CAEEL	38.8%		PNEDRYMKPEIDAVLEMIRENRIWEAVLPHLET					
15	SWALL:Q9HLI6	41.0%		LDHDRPPSFDIETIRKMMDKKEFISALP-----					
16	SWALL:HUTH_MOUSE	38.6%		WIKDRFMAPDIEAAHRLLLLDQKVWEVAAPYIEK					
17	SWALL:BAB29407	38.6%		WIKDRFMAPDIEAAHRLLLLDQKVWEVAAPYIEK					
18	SWALL:HUTH_RAT	38.2%		WIKDRFMAPDIEAAHRLLLLDQKVWEVAAPYIEK					
19	SWALL:AAG53586	39.8%		YDDDRFFAPDIEAAISL NKGSLVGLLPAFL--					
20	SWALL:Q9KKE0	38.9%		PIATIVR-----					
21	SWALL:Q9HQD5	42.2%		PAGDRALADDMAAVGDLVRAGLVEDAVARALDA					

DocId: 3428860



TOE40" 342E662

Figure 14, cont'd.

KEY:

983831 : HAL

1	CAC21618	: Streptomyces coelicolor
2	HUTH_STRGR	: Streptomyces griseus
3	HUTH_DEIRA	: Deinococcus radiodurans
4	BAB16159	: Agrobacterium rhizogenes
5	Q9KWE4	: Agrobacterium rhizogenes
6	HUTH_BACSU	: Bacillus subtilis
7	Q9KSQ4	: Vibrio cholerae
8	Q9HU85	: Pseudomonas aeruginosa
9	Q9KBE6	: Bacillus halodurans
10	HUTH_PSEPU	: Pseudomonas putida
11	HUTH_RHIME	: Rhizobium meliloti
12	Q9HU90	: Pseudomonas aeruginosa
13	HUTH_HUMAN	: Human
14	HUTH_CAEEL	: Caenorhabditis elegans
15	Q9HLI6	: Thermoplasma acidophilum
16	HUTH_MOUSE	: Mouse
17	BAB29407	: Mus musculus (Mouse)
18	HUTH_RAT	: Rat
18	AAG53586	: uncultured bacterium pCosAS1
20	Q9KKE0	: Rhizobium meliloti
21	Q9HQD5	: Halobacterium sp

TEETH "SHED"

Figure 15

STRG	6	VVVGTSGTTAEDVVARHGARVELSAAVEALAAARLIVDALAAKPEPVYGVSTGFGAL	
"HAL"	7	ITLGLSGATADDVIAVARHEARISISPPQVLEELASVRAHIDALASADTPVYGISTGFGAL	
		* * * * *	* * * * *
STRG,	66	ASRHIGTELRAQLQRNIVRSHAAAGMPRVEREVVRALMFLRLKTVASGHTGVRPEVAQTM	
HAL	67	ATHIAPEDRAKLQRLIRSHAAAGMPVEREVVRALMFLRAKTLASGRTGVRPVLVETM	
		* * * * *	* * * * *
STRG	126	ADVLNAGITPVVHEYGSLGCSGDLAPLSHCALTLMGEGEAEGPDGTVRPAGELLAAHGIA	
HAL	127	VGMLNAGITPVVREYGSLGCSGDLAPLSHCALVLMGEGEATDAHGDIRPVPELFAEAGLT	
		* * * * *	* * * * *
STRG	186	PVELREKEGLALLNGTDGMLGMLMALADLRNLYTSADITAAALSLEALLGTDKVLAPELH	
HAL	187	PVELAEKEGLALVNGTDGMLGQLIMALADLDELDDIADATAAMSVEAQLGTDQVFRAELH	
		* * * * *	* * * * *
STRG	246	A-IRPHPGQVSADNMSRVLAGSLTGHHQDDAPRVQDAYSVRCAPQVNGAGRDTLDHAA	
HAL	247	EPLRPHPGQGRSAQNMFAFLADSPIVASHREGDGRVQDAYSLRCSPQVTGAARDTIAHAR	
		* * * * *	* * * * *
STRG	305	LVAGRELASSVDNPVLPDGRVESNGNFHGAPVAYVLDFLAIVAADLGSICERRTDRLD	
HAL	307	LVATRELAAAIIDNPVLPSPGEVTSNGNFHGAPVAYVLDFLAIAVADLGSIAERRTDRMLD	
		* * * * *	* * * * *

TOE TTD" 344E3B3

Figure 15, Cont'd.

STRG	365	KNRSHGLPPFLADDAGVDSGLMIAQYTQAALVSEMKRRLAVPASADSI	IPSSAMQEDHVS	MG
HAL	367	PARSRDLPAFLADDPGVDSGMMIAQYTQAGLVAENKRRLAVPASVDSI	IPSSAMQEDHVS	LG
		** ** *	** ** *	** ** *
STRG	425	WSAARKLRTAVDNLARI VAVELYAATRAIELRAAEG	LT	PAPASEAVVAALRAAG
HAL	427	WHAARKLRTSVANLRRILAVEMLIAGRALDLRAP--	LKPGPATGAVLE	VLRSKVA-GPGQ
		* ** *	* ** *	* ** *
STRG	485	DRFLAPDLAAADTFVREGRLVAAVE		
HAL	484	DRFLSAELEAAAYDLLANGSVHKALE		
		***	* ** *	* ** *